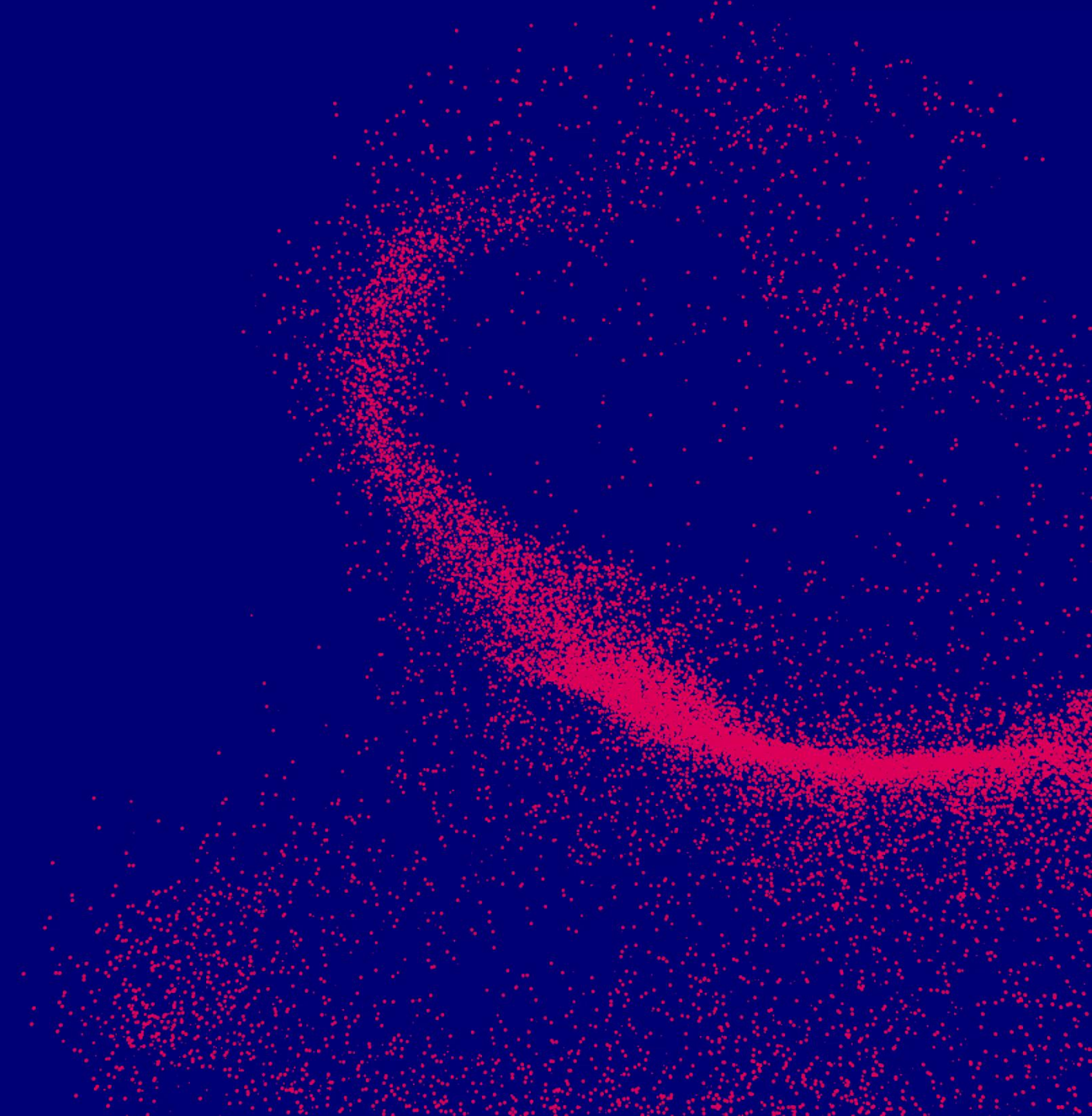


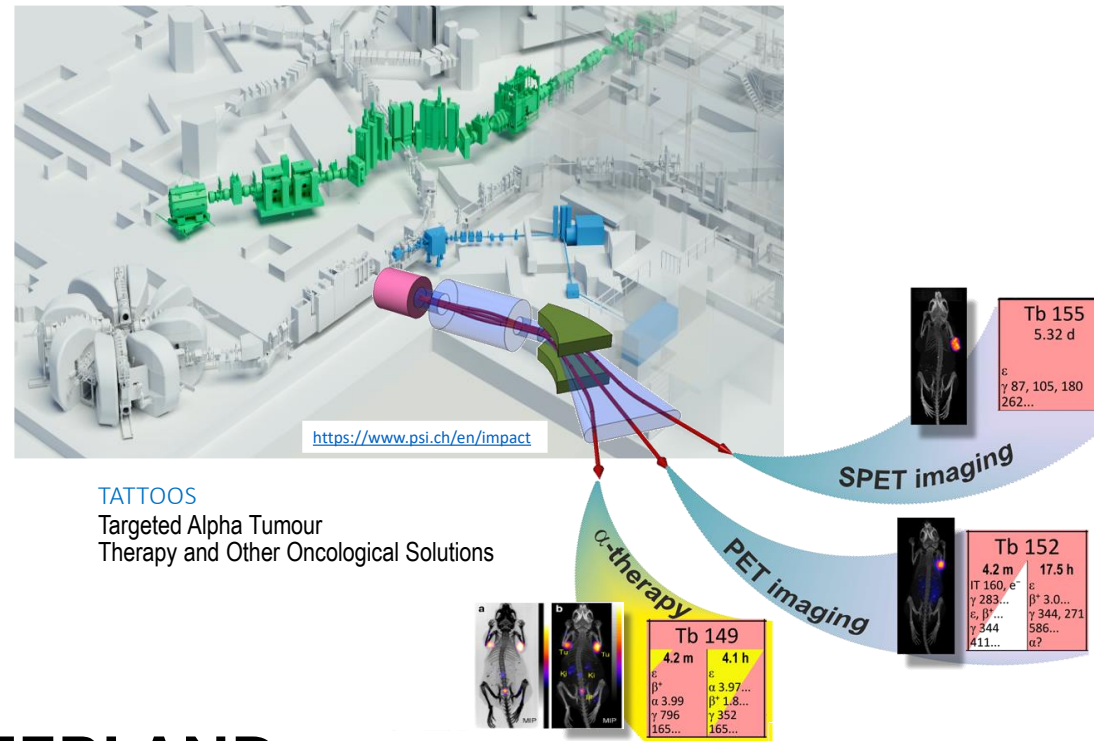
TATOOS





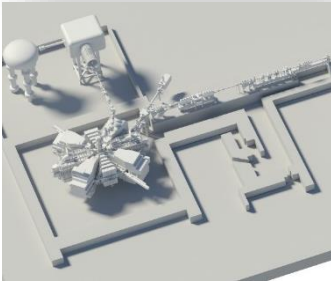
IMPACT-TATTOOS

Novel radionuclide production in SWITZERLAND: a design status report



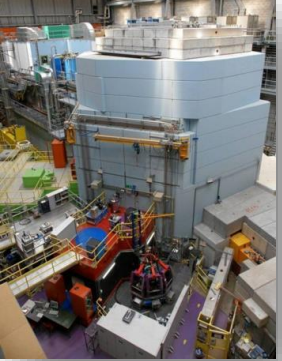
Robert Eichler :: Laboratory of Radiochemistry & University of Bern for the IMPACT collaboration
PRISMAP Radiolanthanide Workshop August 3 – August 6 2024, Villigen , PSI, Switzerland

“Terbium Sisters” for Theragnostics: Terbium A Multi-Purpose Element



70 MeV PROTONS

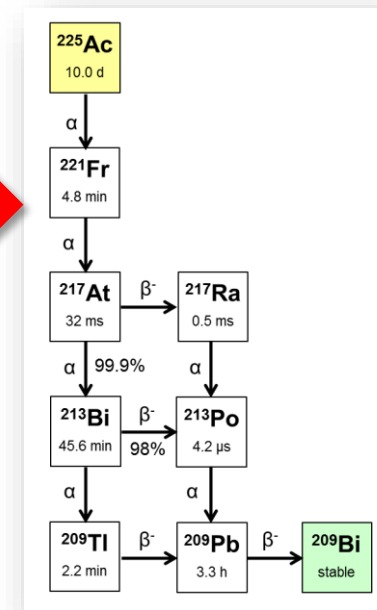
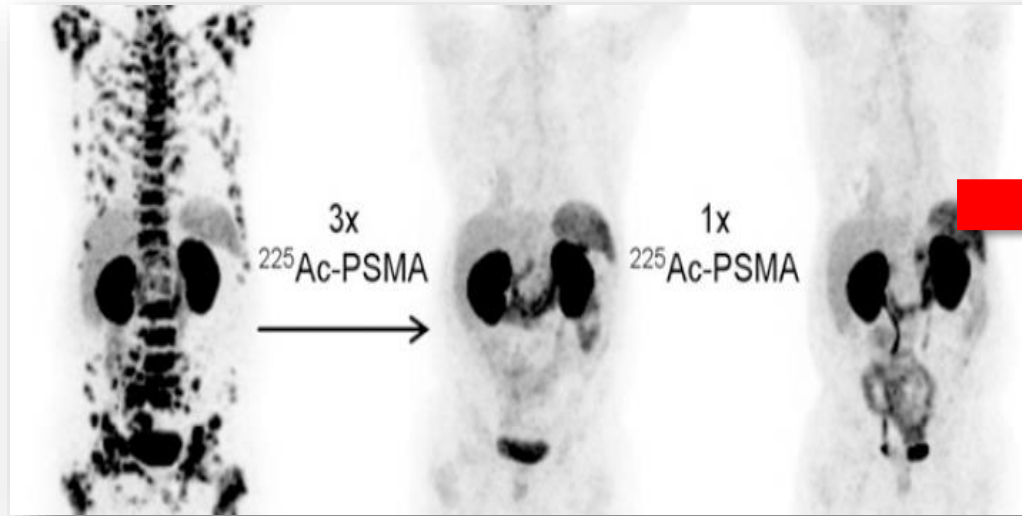
Isotope	T _{1/2}	Decay mode	Potential use
149Tb	4.1 h	α (3.6 MeV); β ⁺ (800 keV)	α-particle therapy
152Tb	17.5 h	β ⁺ (1275 keV)	PET
155Tb	5.32 d	γ (87, 105 ... keV)	SPECT
161Tb	6.88 d	β ⁻ (593 keV)	β ⁻ -particle & Auger electron therapy



Dy 150 7.2 m	Dy 151 17 m	Dy 152 2.4 h	Dy 153 6.29 h	Dy 154 3.0 · 10 ⁹ a	Dy 155 10.0 h	Dy 156 0.056	Dy 157 8.1 h	Dy 158 0.095	Dy 159 144.4 d	Dy 160 2.329	Dy 161 18.889	Dy 162 25.475	Dy 163 24.8
Tb 149 4.2 m	Tb 150 4.1 h	Tb 151 3.67 h	Tb 152 17.5 h	Tb 153 2.34 d	Tb 154 23 h	Tb 155 5.32 d	Tb 156 5.4 h	Tb 157 99 a	Tb 158 10.5 s	Tb 159 100	Tb 160 72.3 d	Tb 161 6.90 d	Tb 162 7.76 m
Gd 148 74.6 a	Gd 149 9.28 d	Gd 150 1.8 · 10 ⁹ a	Gd 151 120 d	Gd 152 1.1 · 10 ¹⁴ a	Gd 153 239.47 d	Gd 154 2.18	Gd 155 14.80	Gd 156 20.47	Gd 157 15.65	Gd 158 24.84	Gd 159 18.48 h	Gd 160 21.86	Gd 161 3.66 m

NEUTRONS

Targeted Alpha Therapy is a Hot Topic!



Kidney
& Salivary glands



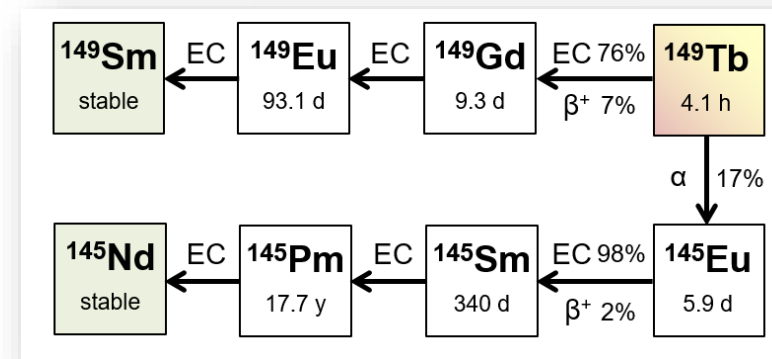
Multiple α/β^- -decaying daughter nuclides can cause much damage,
Resulting in side effects!

4h half-life / alpha & beta+ decay

Transport loss
between Geneva and PSI:
50% \rightarrow ~200 MBq
 \rightarrow barely enough for
preclinical (animal) studies

Tb 149	
4.2 m	4.1 h
ϵ	ϵ
β^+	α 3.97...
α 3.99	β^+ 1.8...
γ 796	γ 352
165...	165...

^{149}Tb has NO α -daughters!



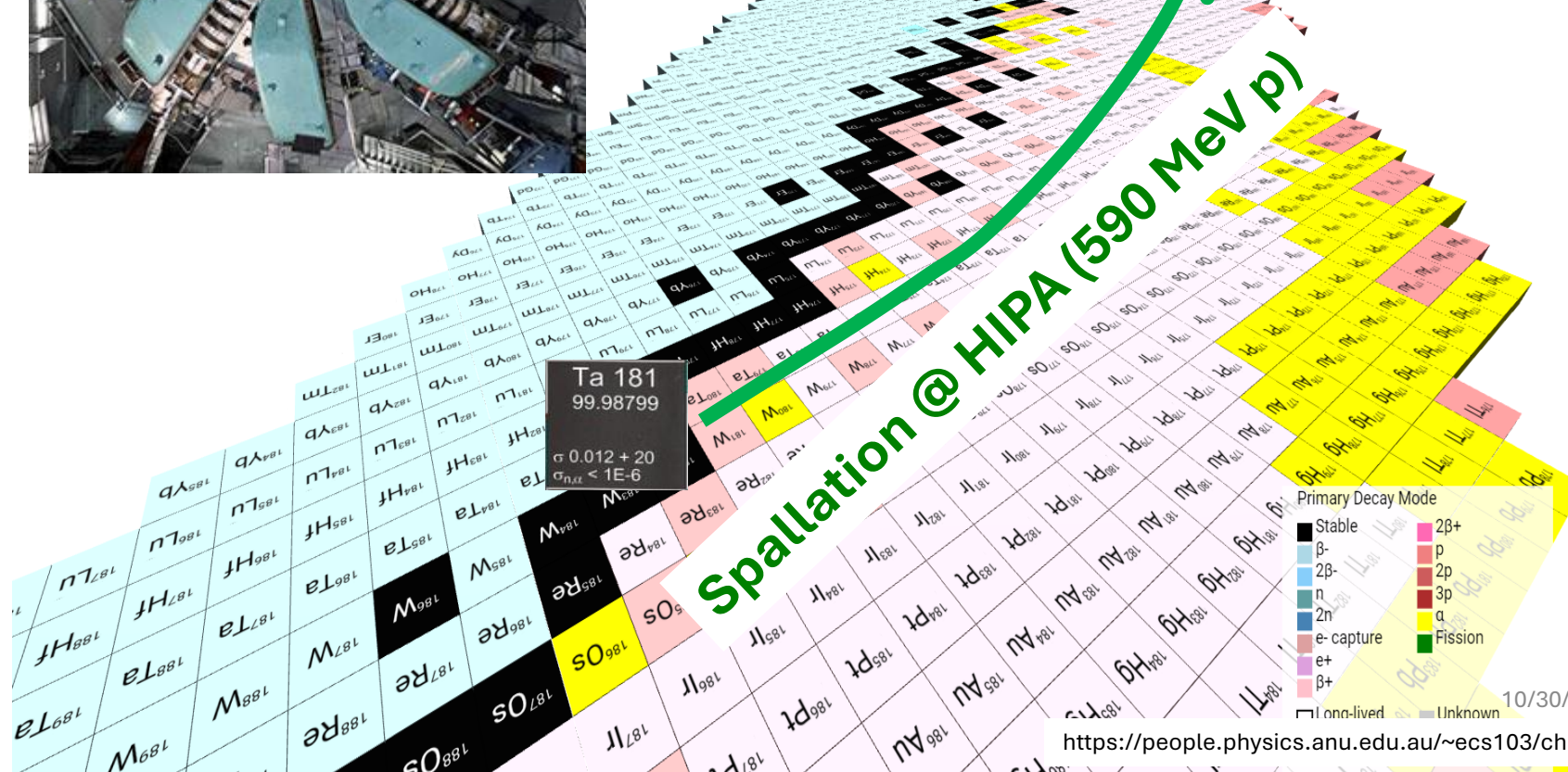
^{149}Tb Production @ PSI-HIPA



World's most powerful DC beam of 2.2 mA 590 MeV protons @ PSI

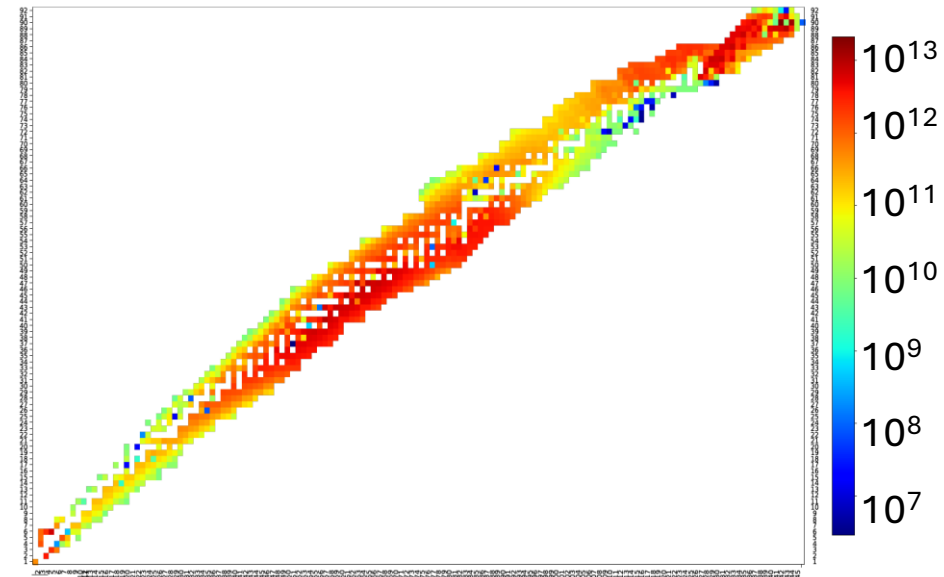
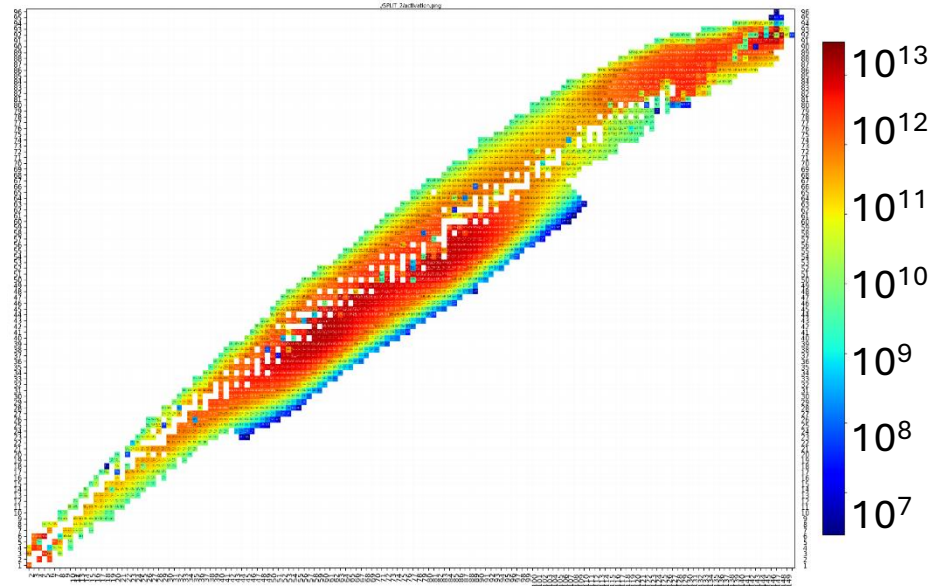
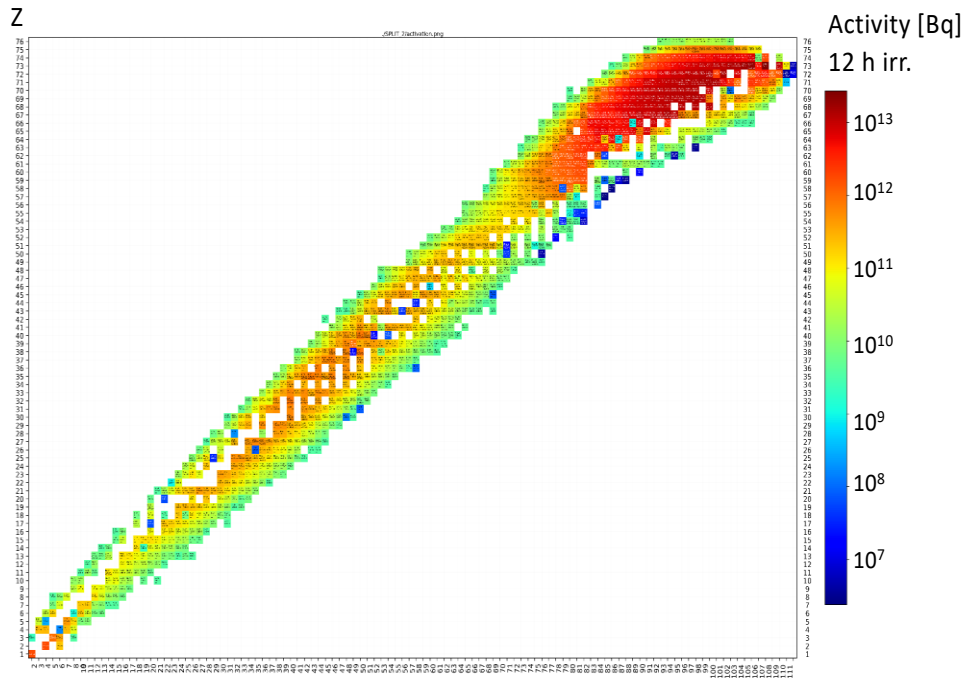


Tb 152		Tb 149	
4.2 m	17.5 h	4.2 m	4.1 h
γ 283;	ϵ	ϵ	ϵ
160...	β^+ 2.8...	β^+	α 3.97
$\epsilon; \beta^+$...	γ 344;	α 3.99	β^+ 1.8
γ 344;	586;	γ 796;	γ 352;
411...	271...	85...	165...



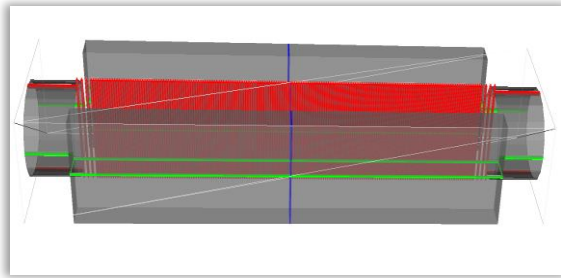
Production of Radionuclides in Spallation

Tantalum 12 h irradiation

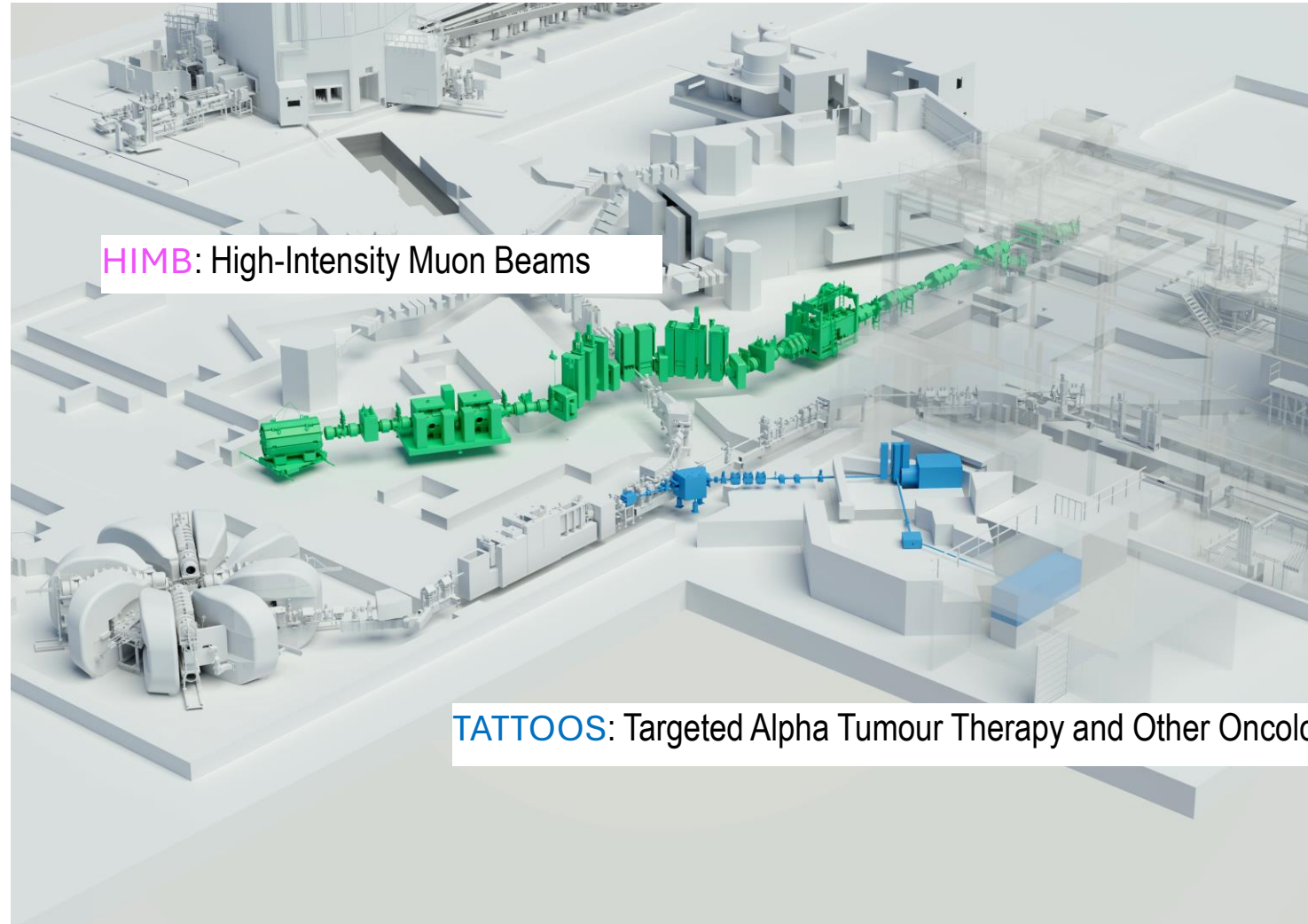
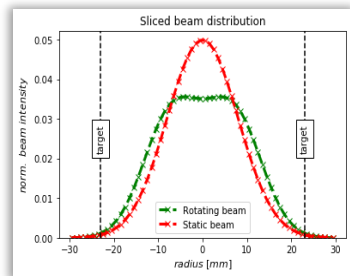


The IMPACT Project 2021

Beam splitter for 100 μ A 590 MeV p



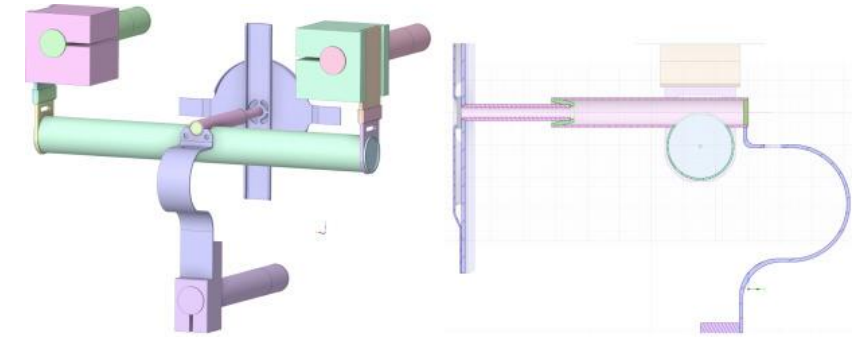
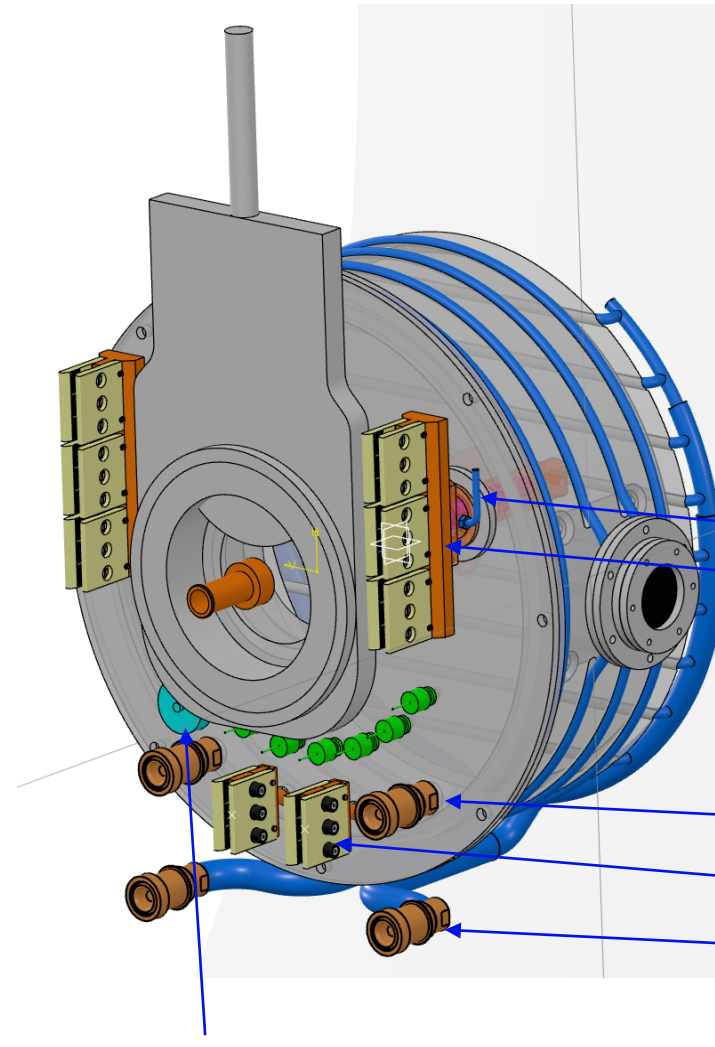
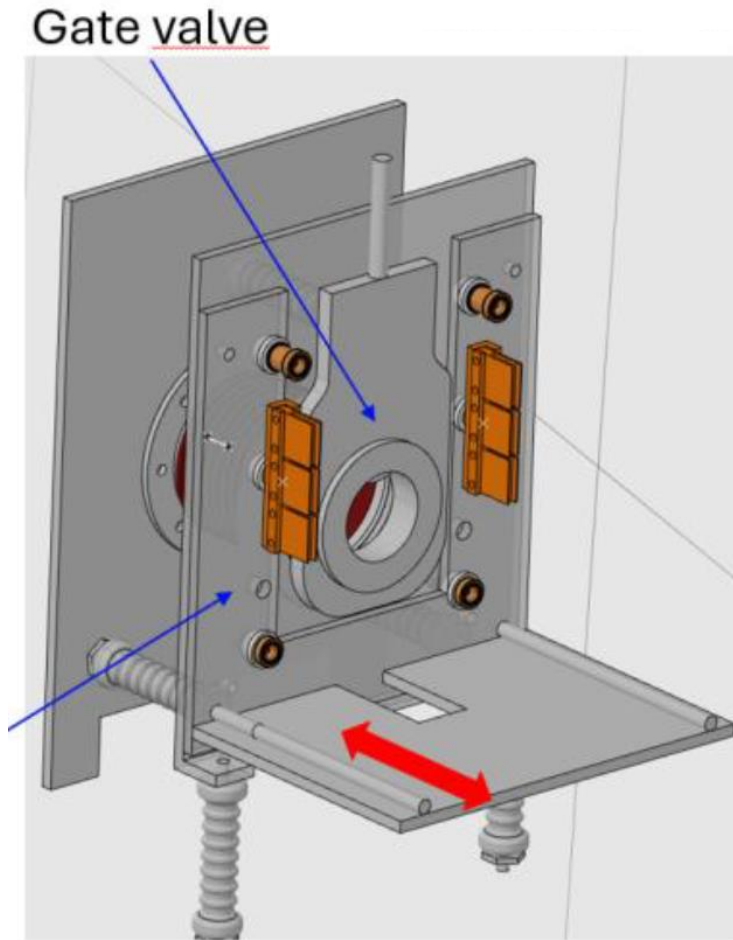
Beam wobbler



HIMB: High-Intensity Muon Beams

TATTOOS: Targeted Alpha Tumour Therapy and Other Oncological Solutions

Target, target chamber & media supply (prelim. concept)



ISOLDE-type target

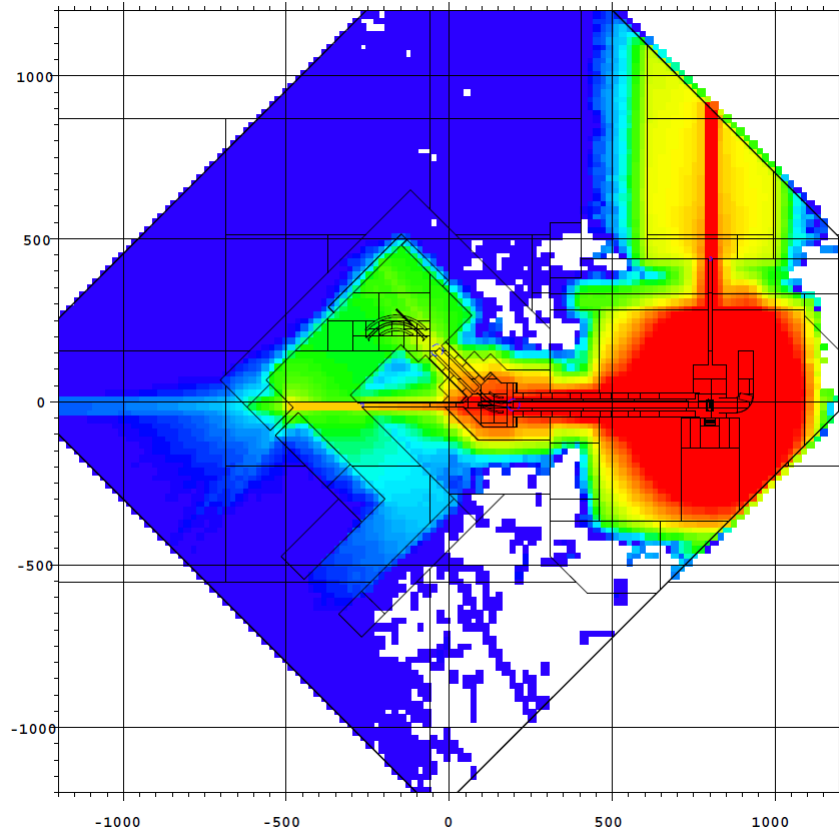
- Cooling of contacts
- High current connector (4'000 A)
- Cooling of media plate
- 2x 1000 A Ionizer heating
- Cooling of chamber

Gas line ioniser (plasma)

In-situ Dose & Shielding Target



Neutrons

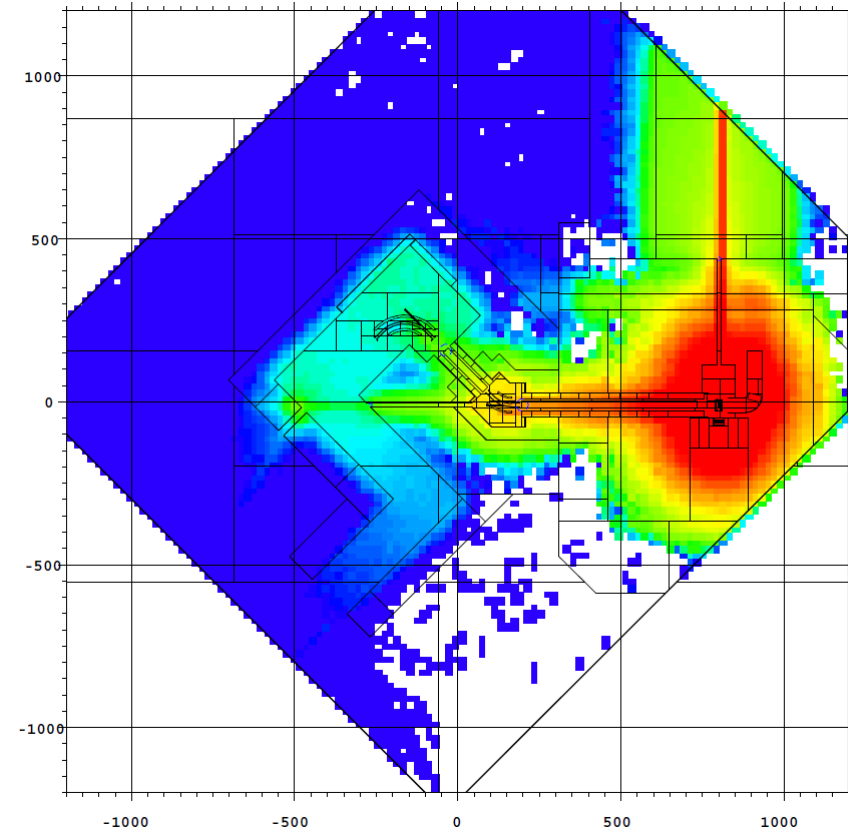


Sv/h



- 1.000E+01
- 1.668E+00
- 2.783E-01
- 4.642E-02
- 7.743E-03
- 1.292E-03
- 2.154E-04
- 3.594E-05
- 5.995E-06
- 1.000E-06

Gamma

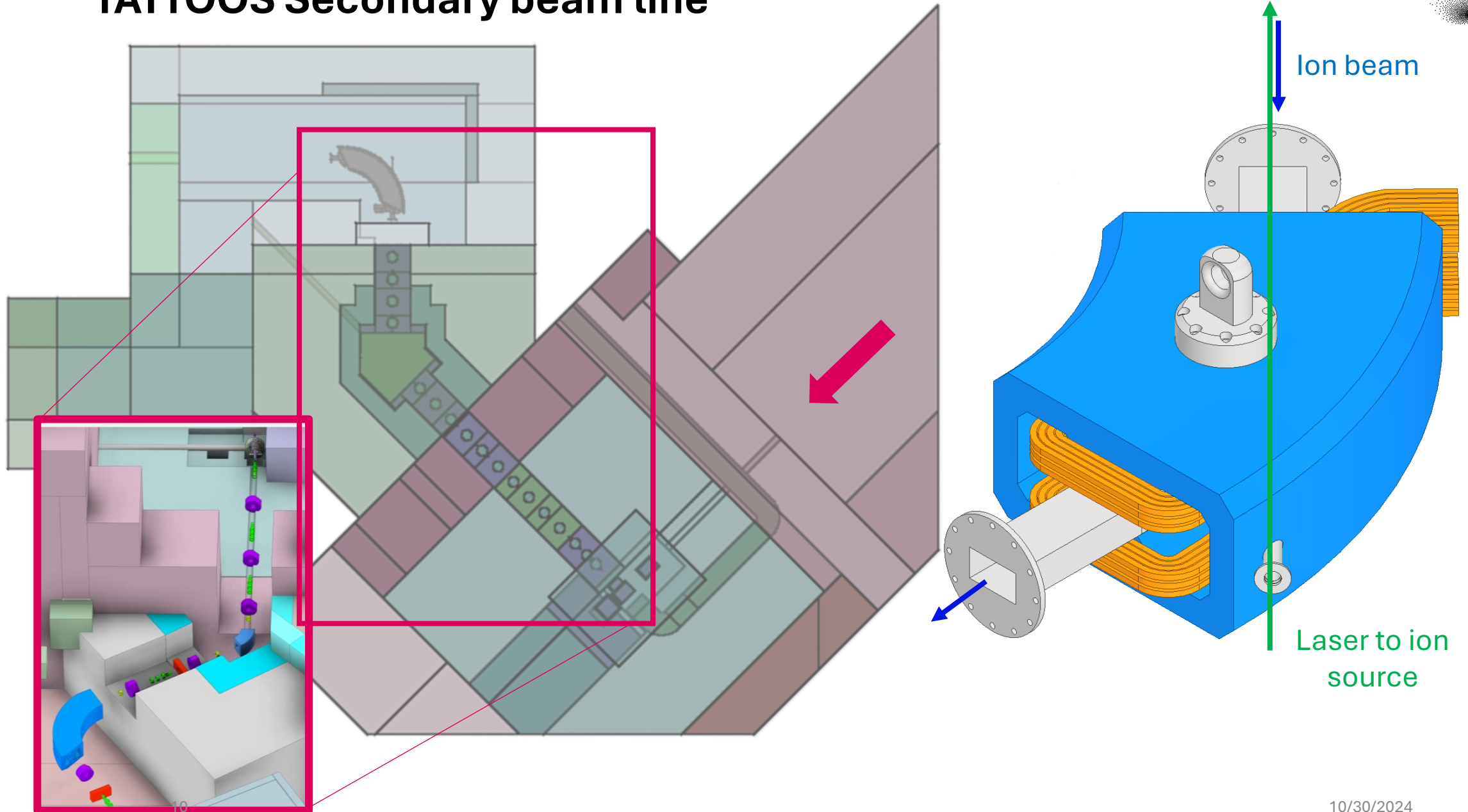


Sv/h

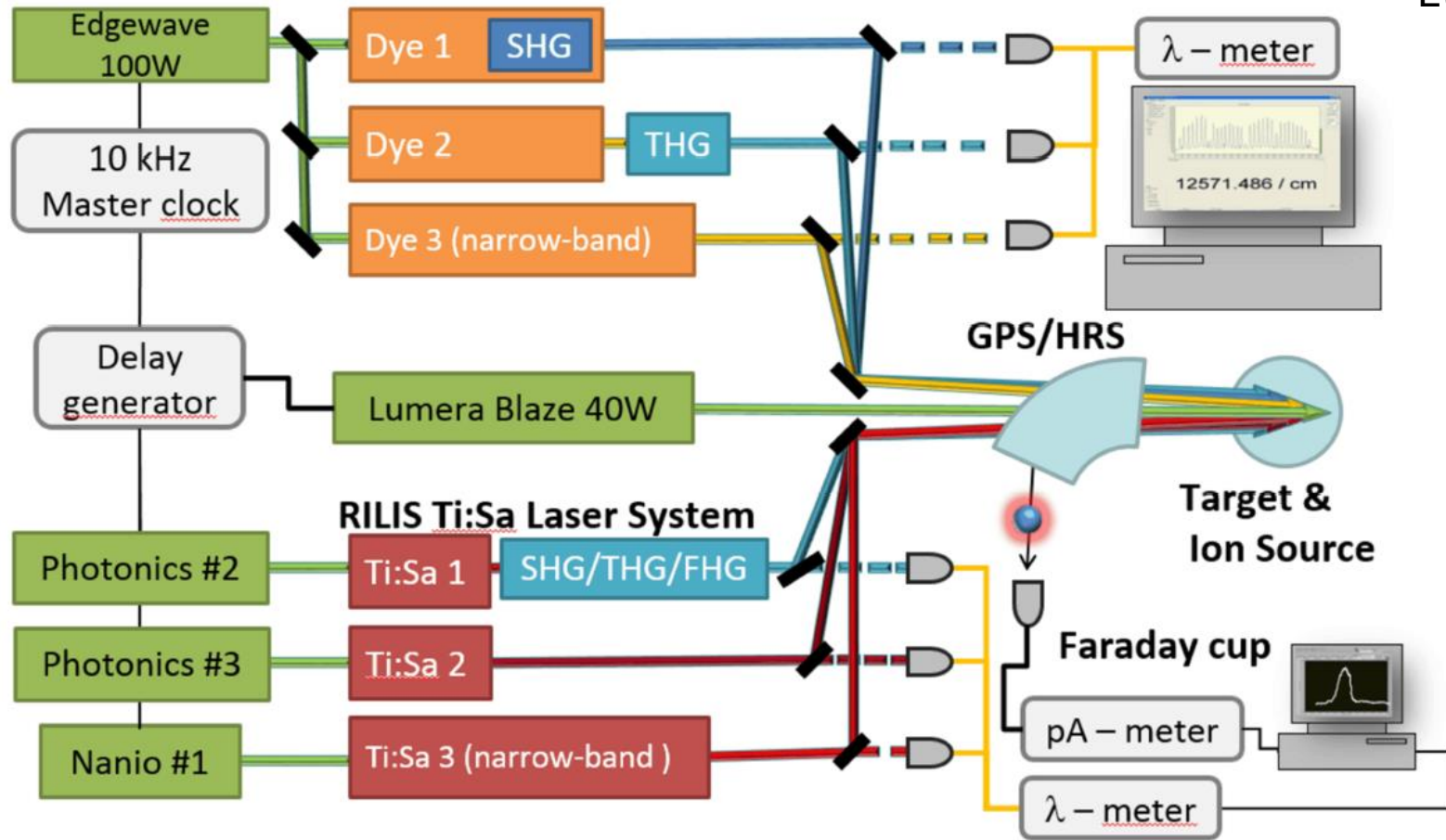


- 1.000E+01
- 1.292E+00
- 1.668E-01
- 2.154E-02
- 2.783E-03
- 3.594E-04
- 4.642E-05
- 5.995E-06
- 7.743E-07
- 1.000E-07

TATTOOS Secondary beam line



Ionization Laser Resonance (RILIS)



Element specific:

Phys. Scr. 85 (2012) 058104

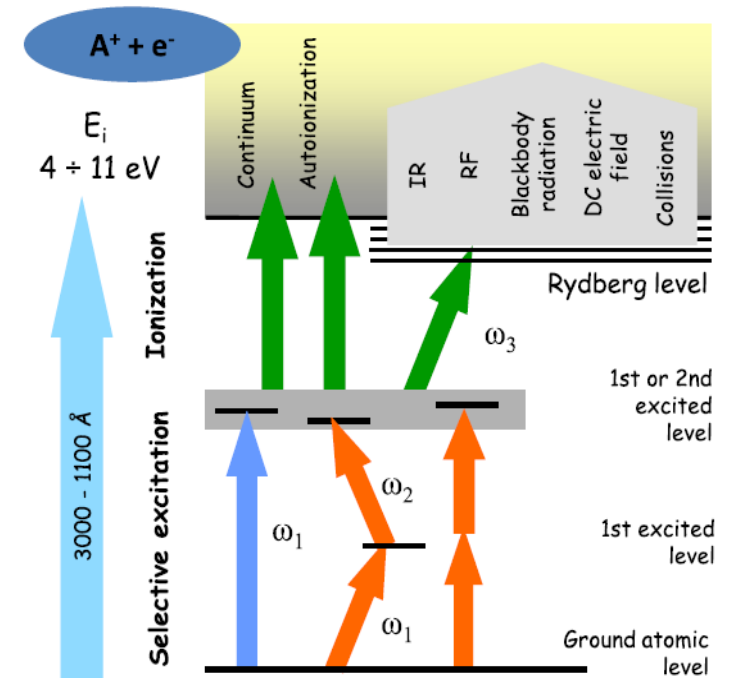
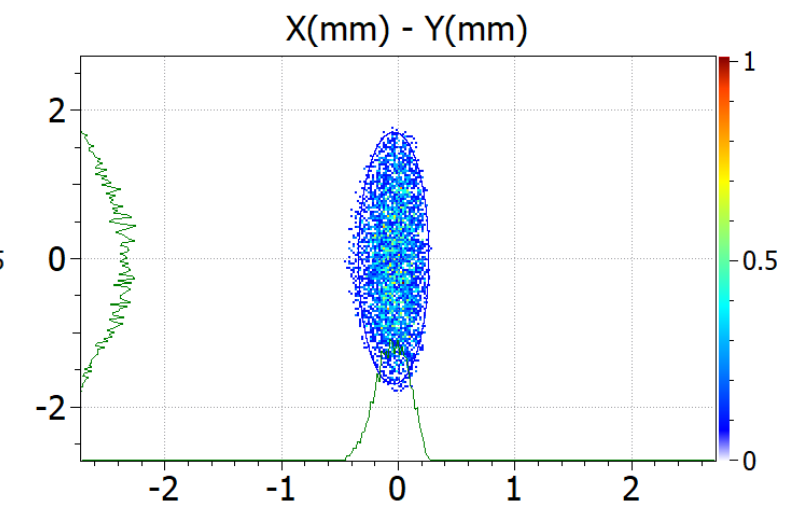
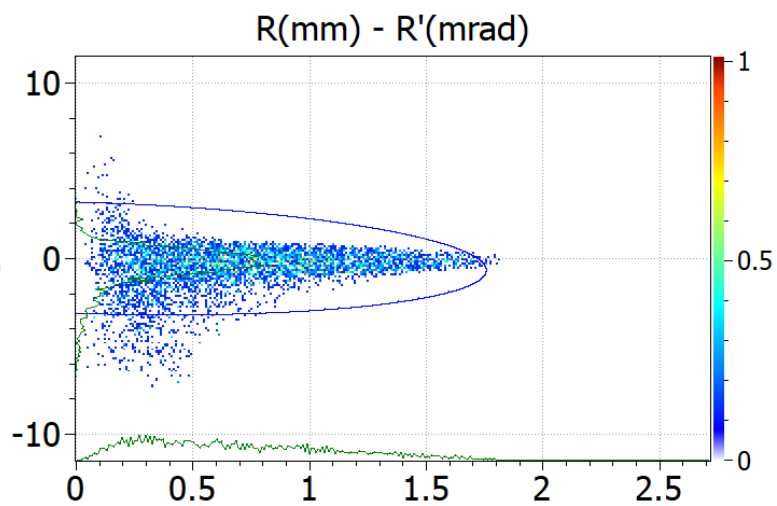
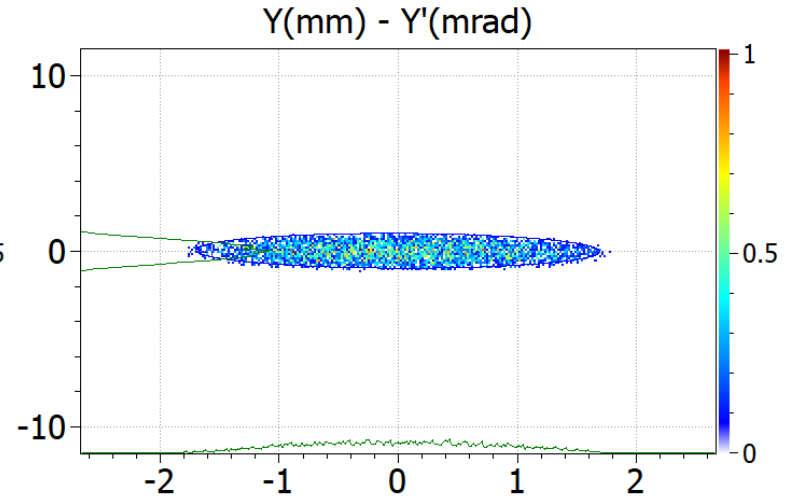
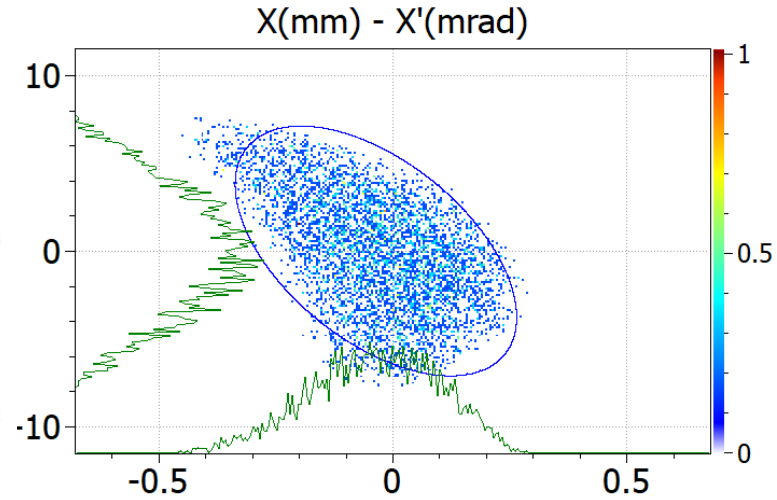
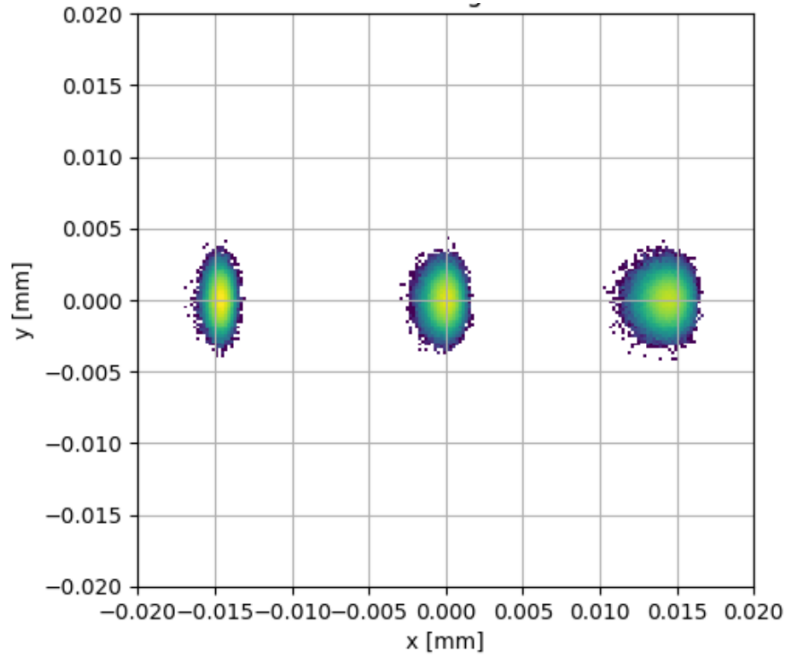
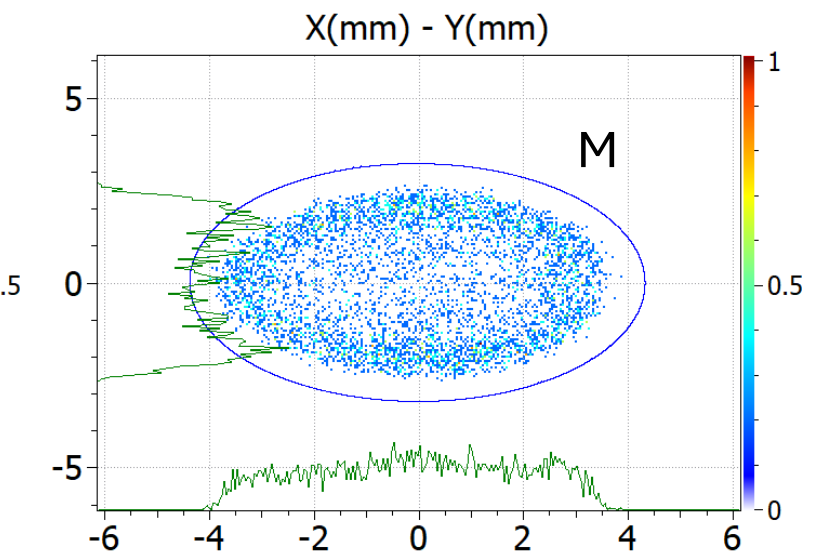
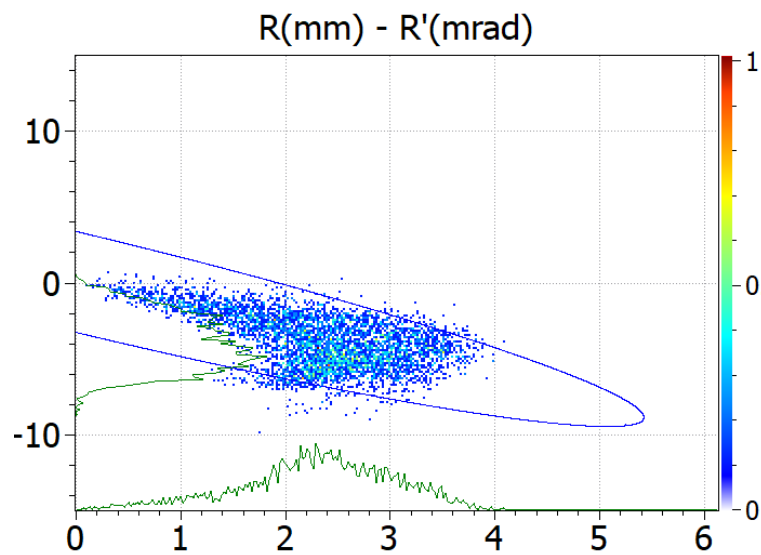
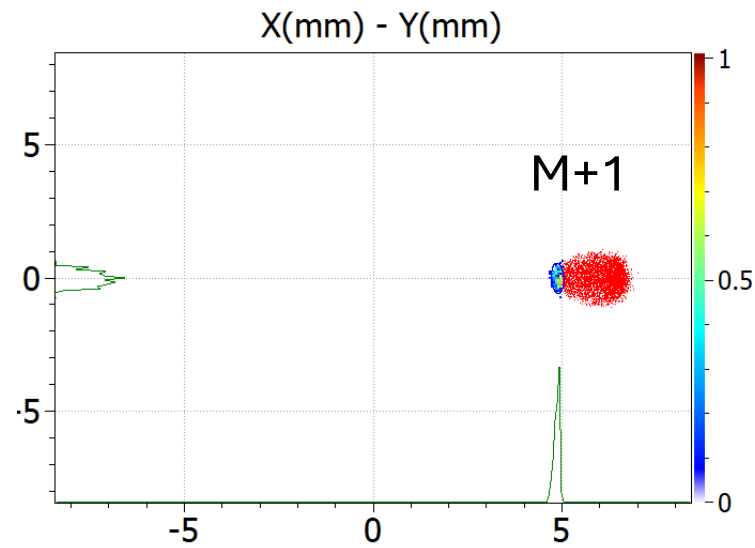
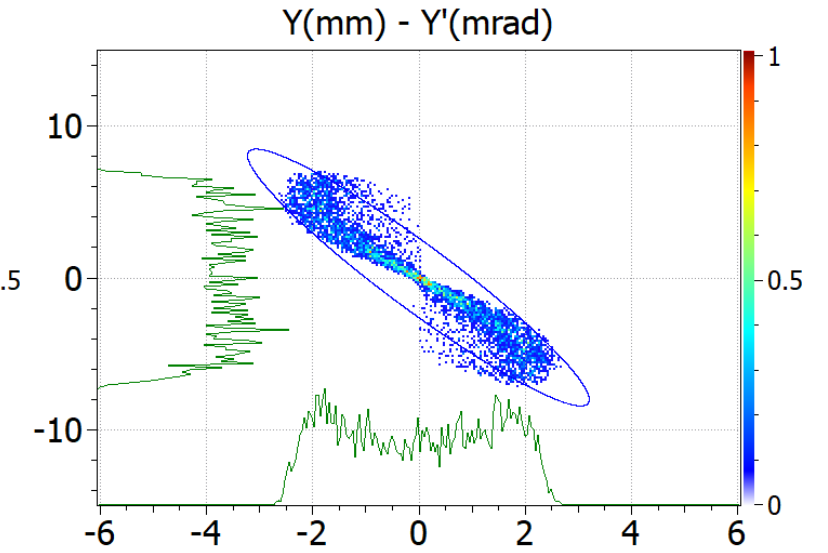
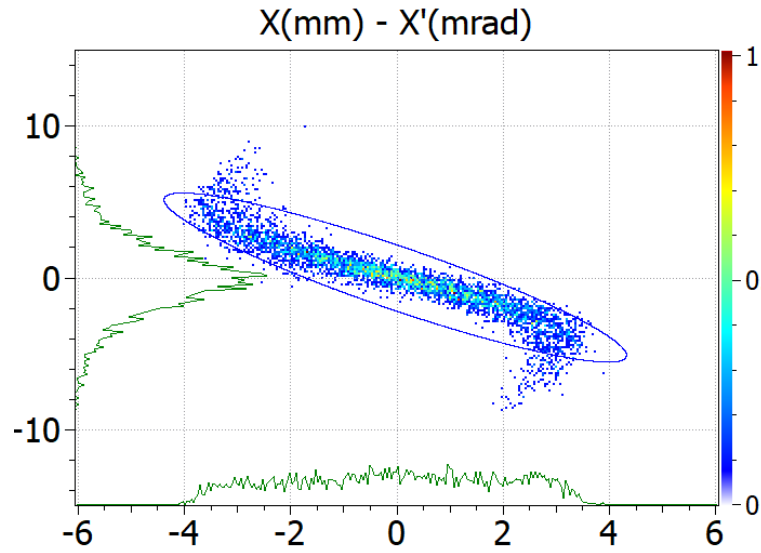
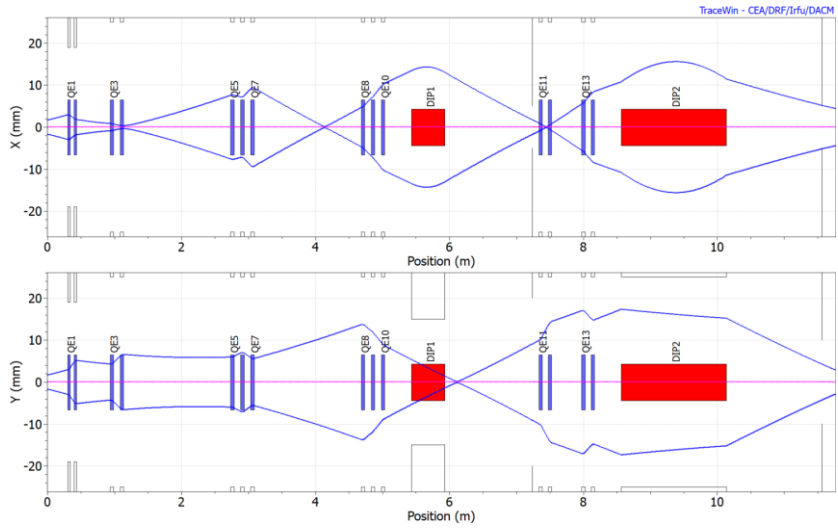


Figure 1. Schemes of resonance ionization.

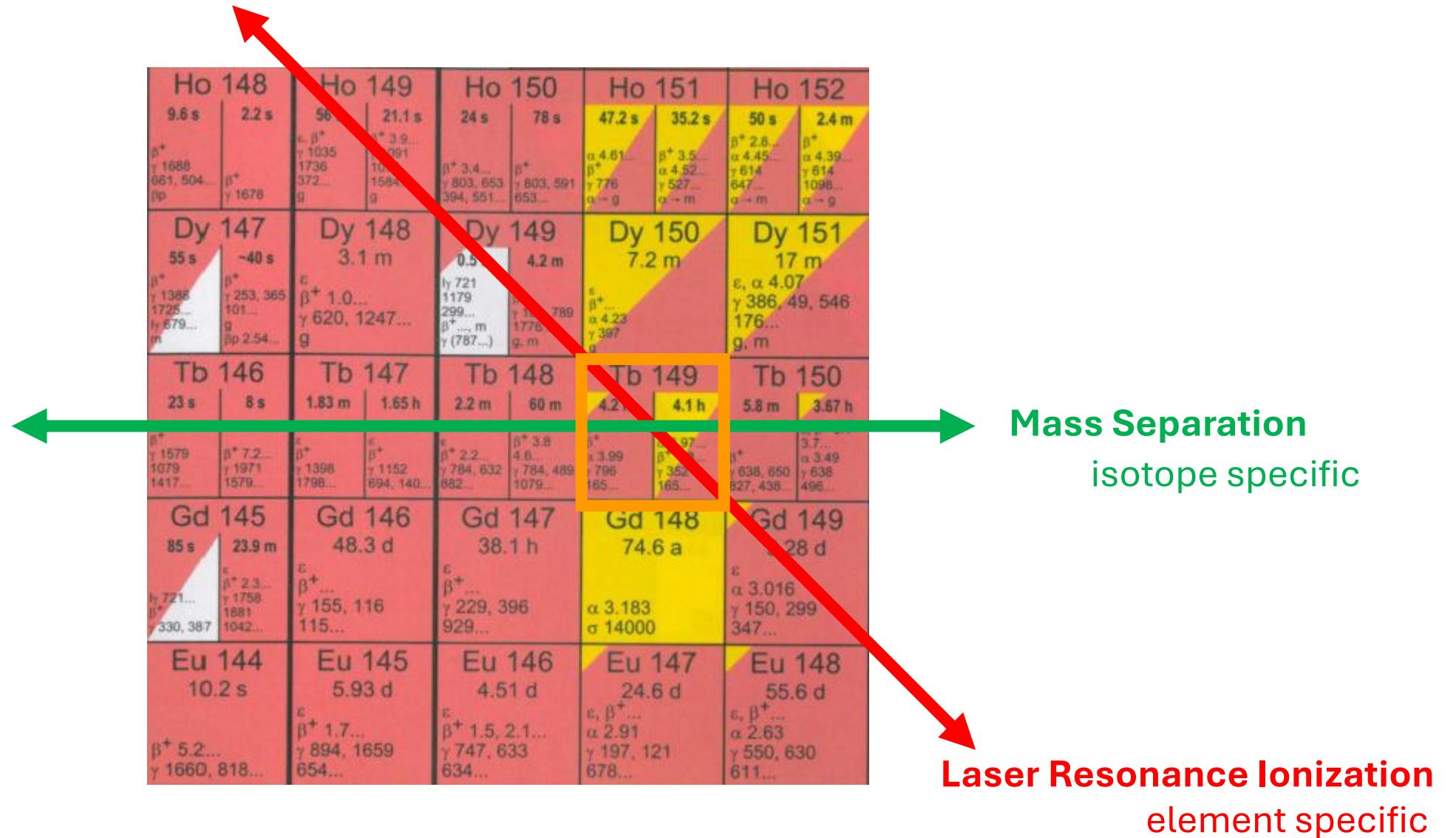
Mass separation simulation @ ion beam of 0.1 uA



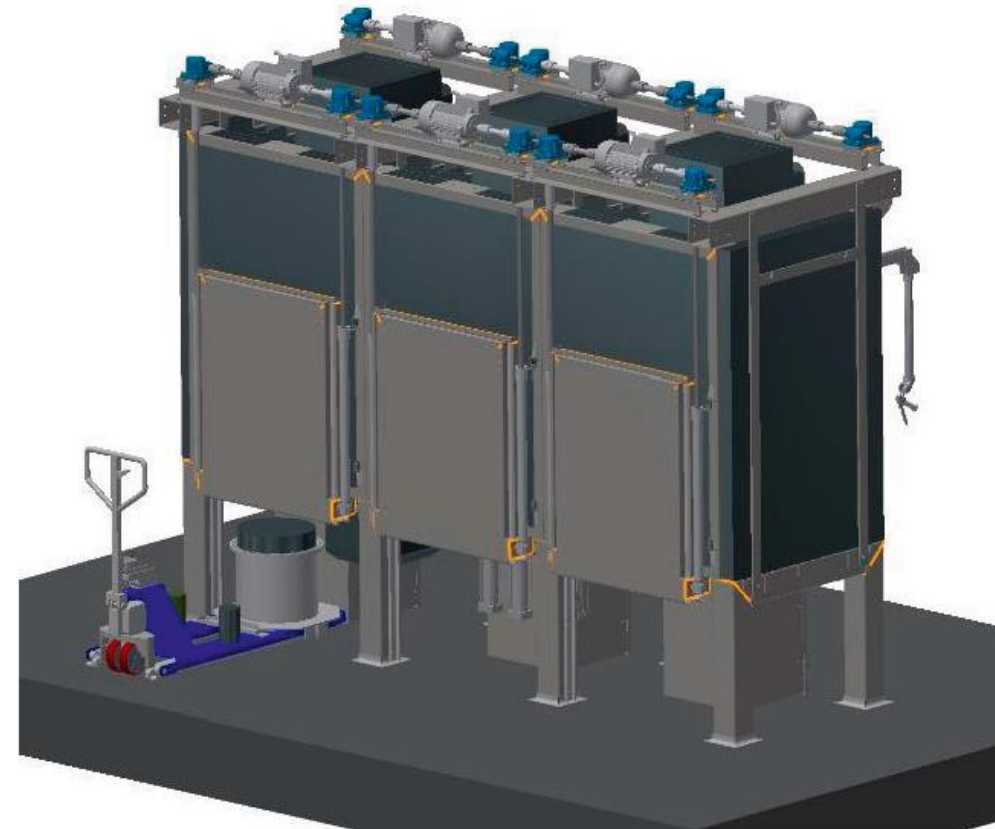
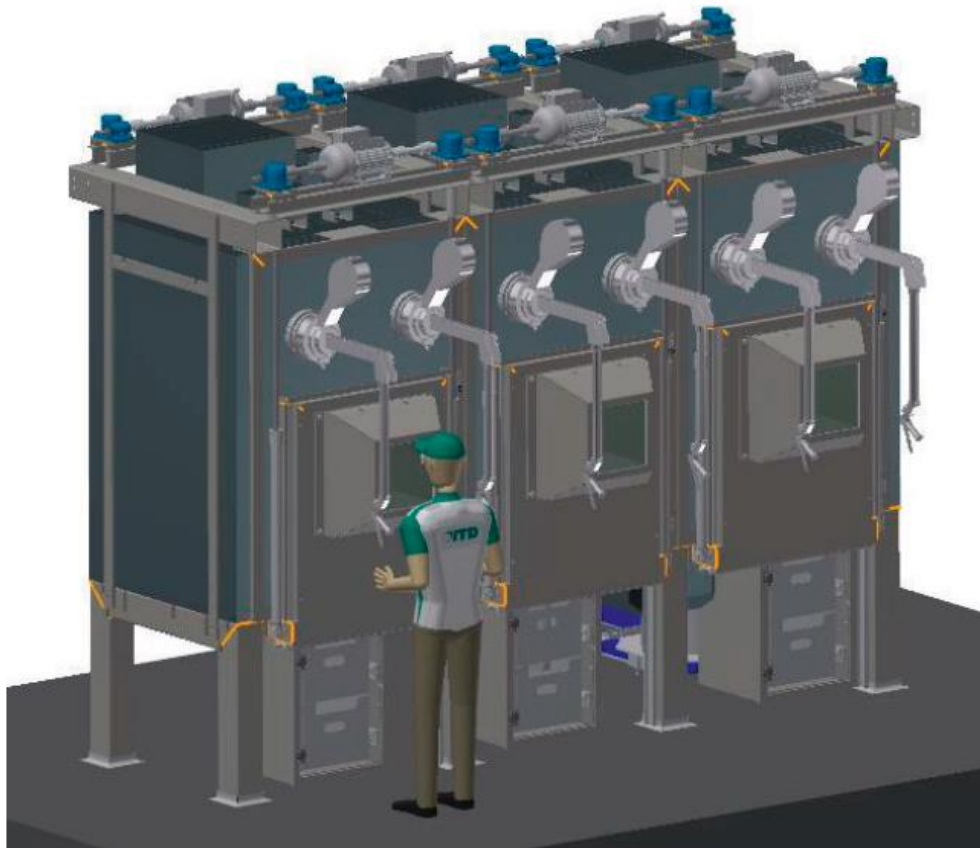
Mass separation simulation @ ion beam of 10-20 μA



The ISOL – principle e.g. Terbium-149



Ion sampling and radiochemical separation of mass isobars in shielded cells



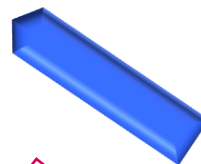
Copyright ITD Dresden

The integrated radiopharmaceutical approach @ PSI through a joint effort

Center for Nuclear Engineering and Sciences (NES)

Laboratory of Radiochemistry

*Expertise in Isotope production,
chemical separation and handling,
fundamental radiochemical research*



Center for Life Sciences (CLS)

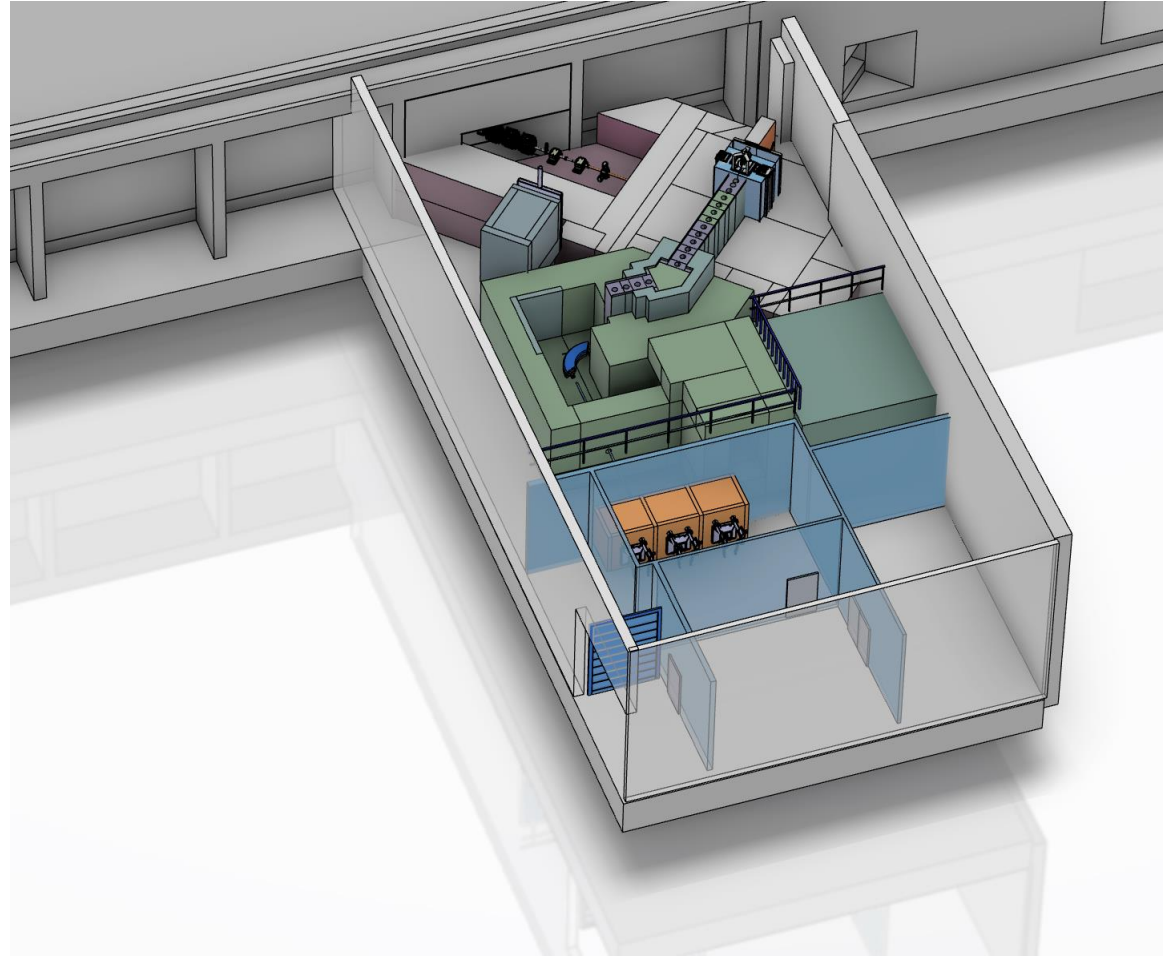
Center for Radiopharmaceutical Sciences

*Expertise in Radiolabeling chemistry,
Preclinical assessment
and Radionuclide therapy development, GMP*

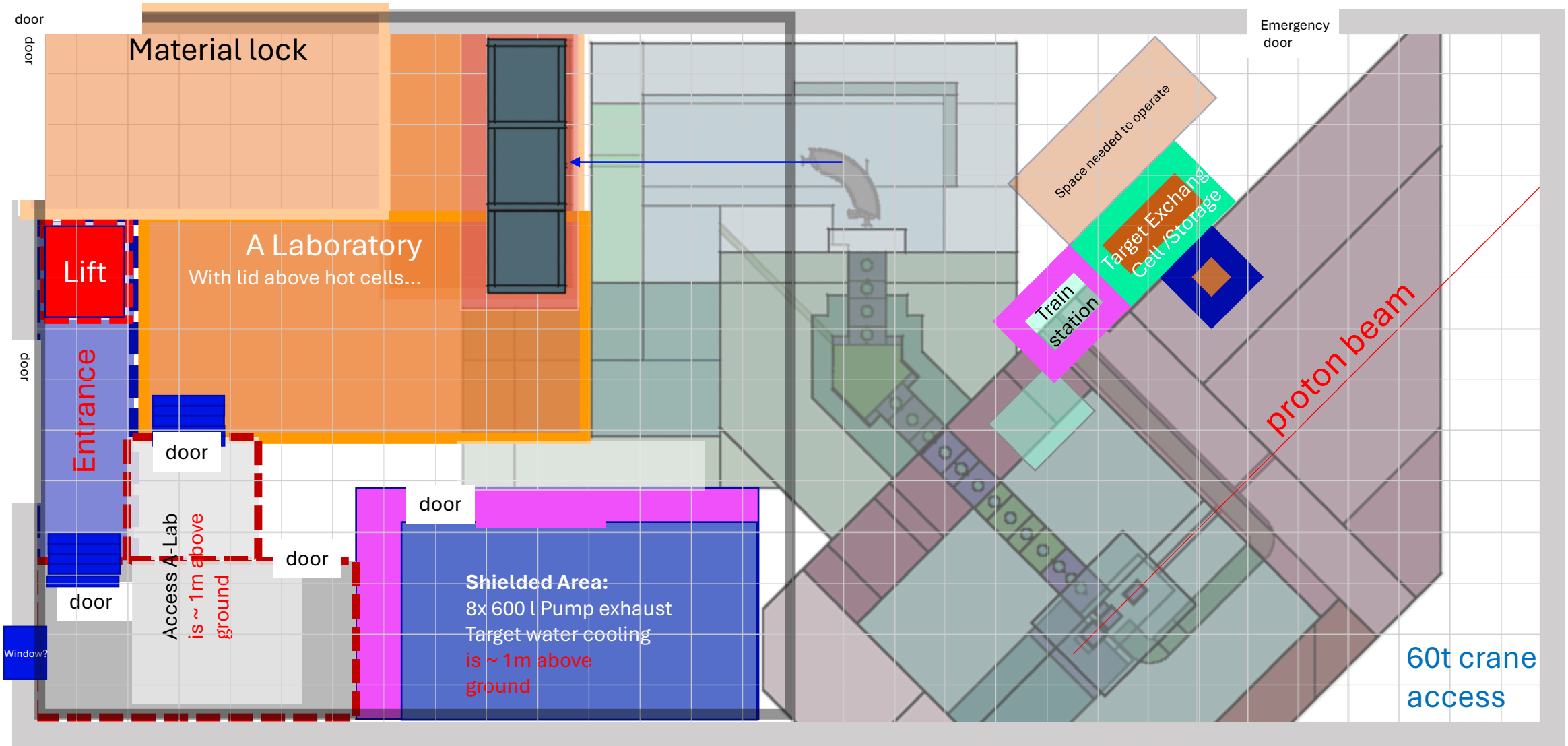



Radionuclide Development group

TATTOOS Building

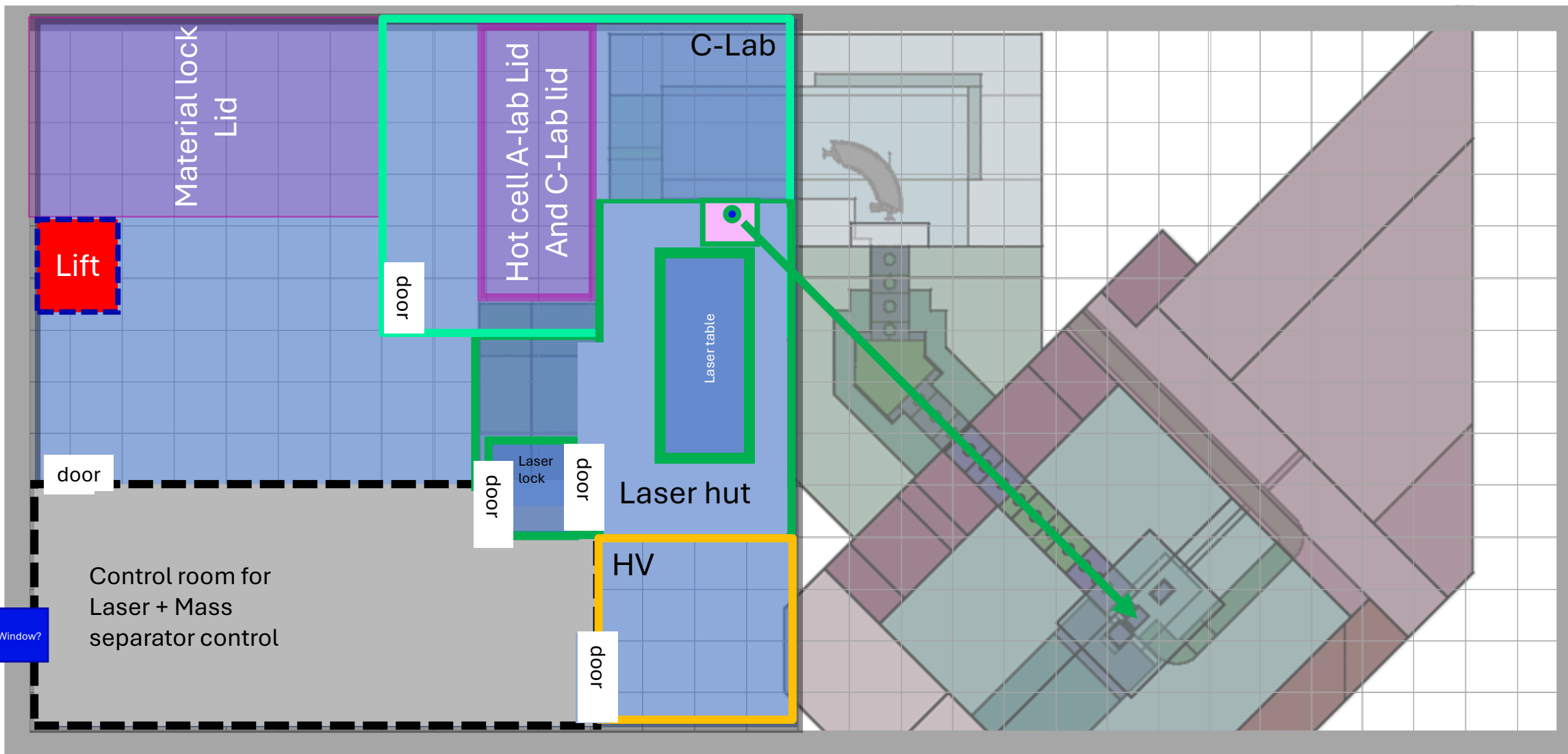


Ground floor

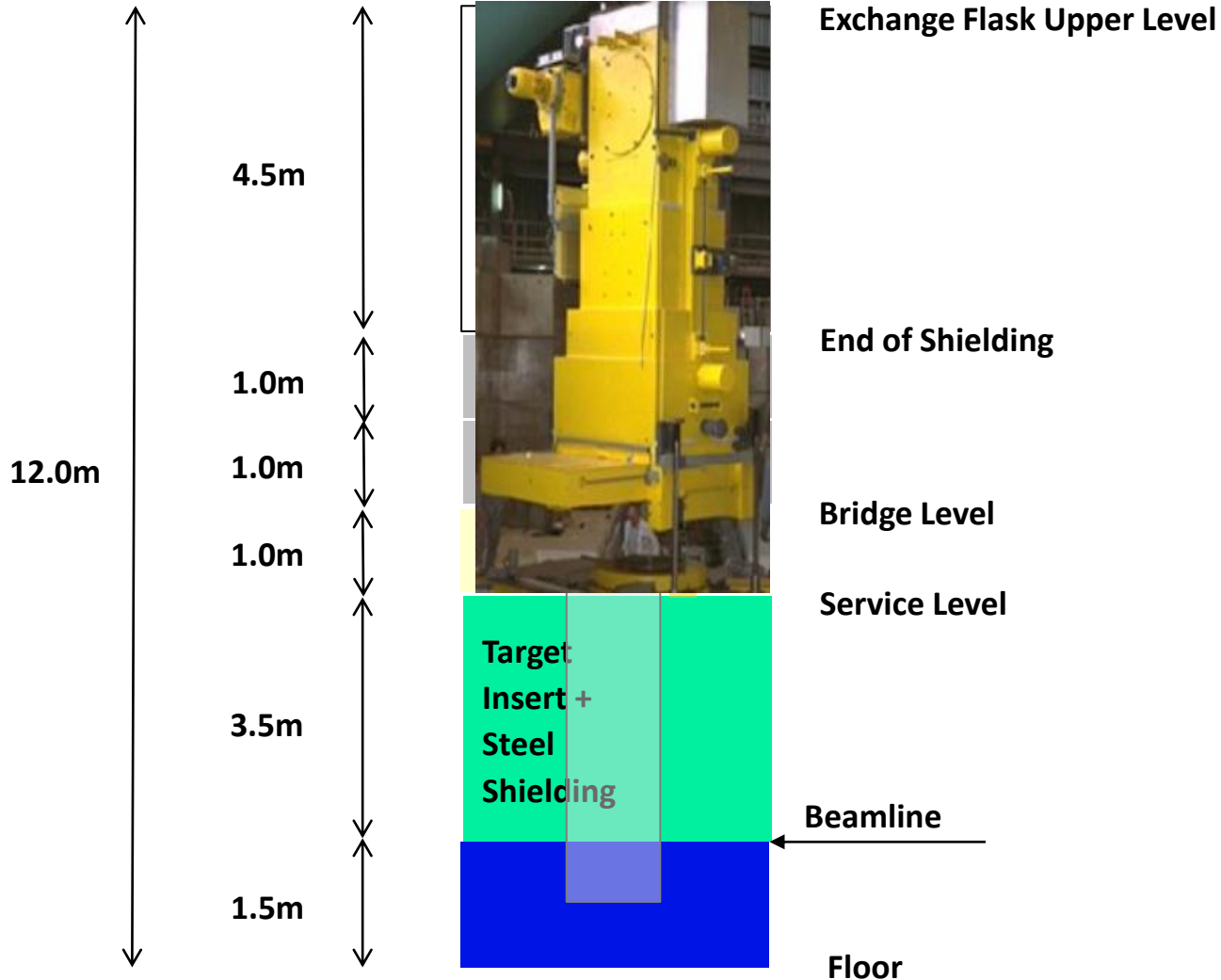



1x1 m²

Second floor on ~5m height

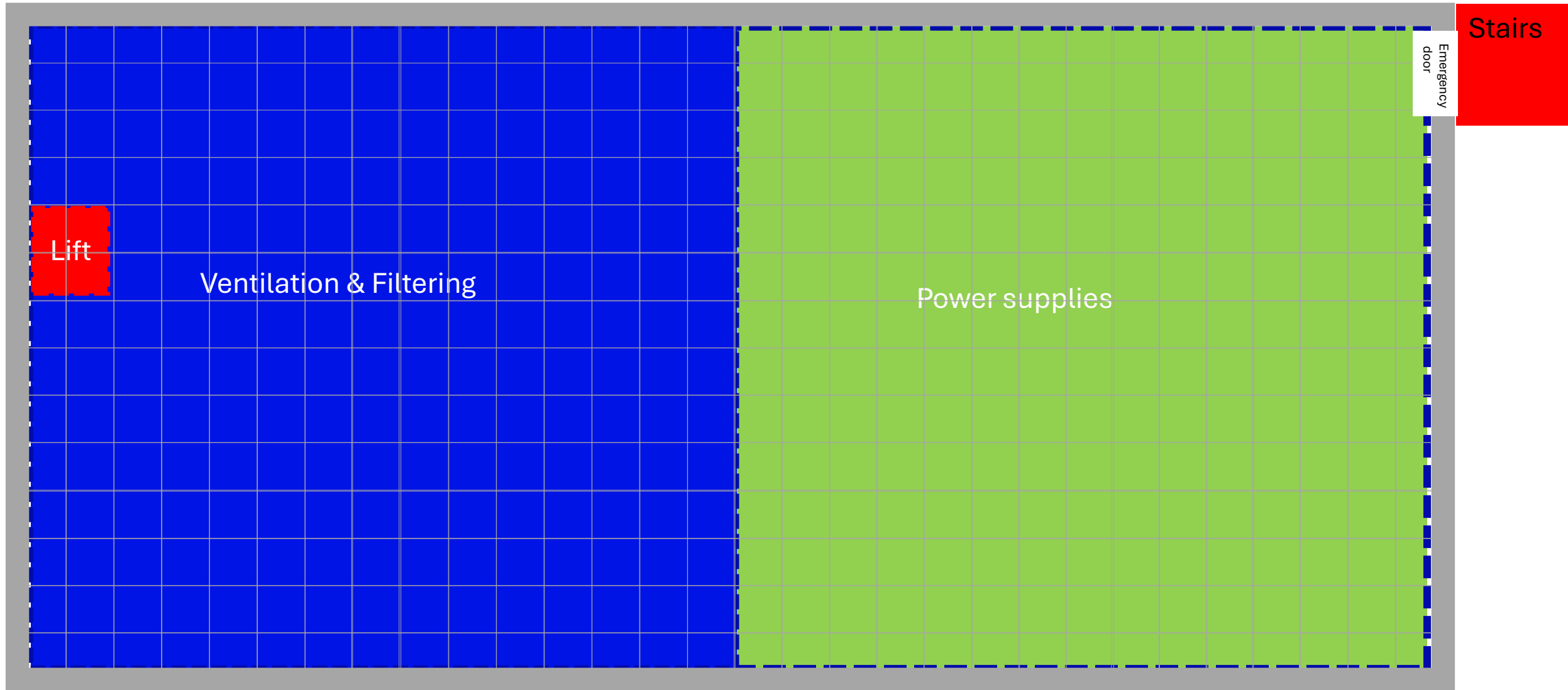


Required TATTOOS Building Height



Full Building:
14 m + Crane (60t) + Roof ≈ **20 m**

Rooftop



Lift

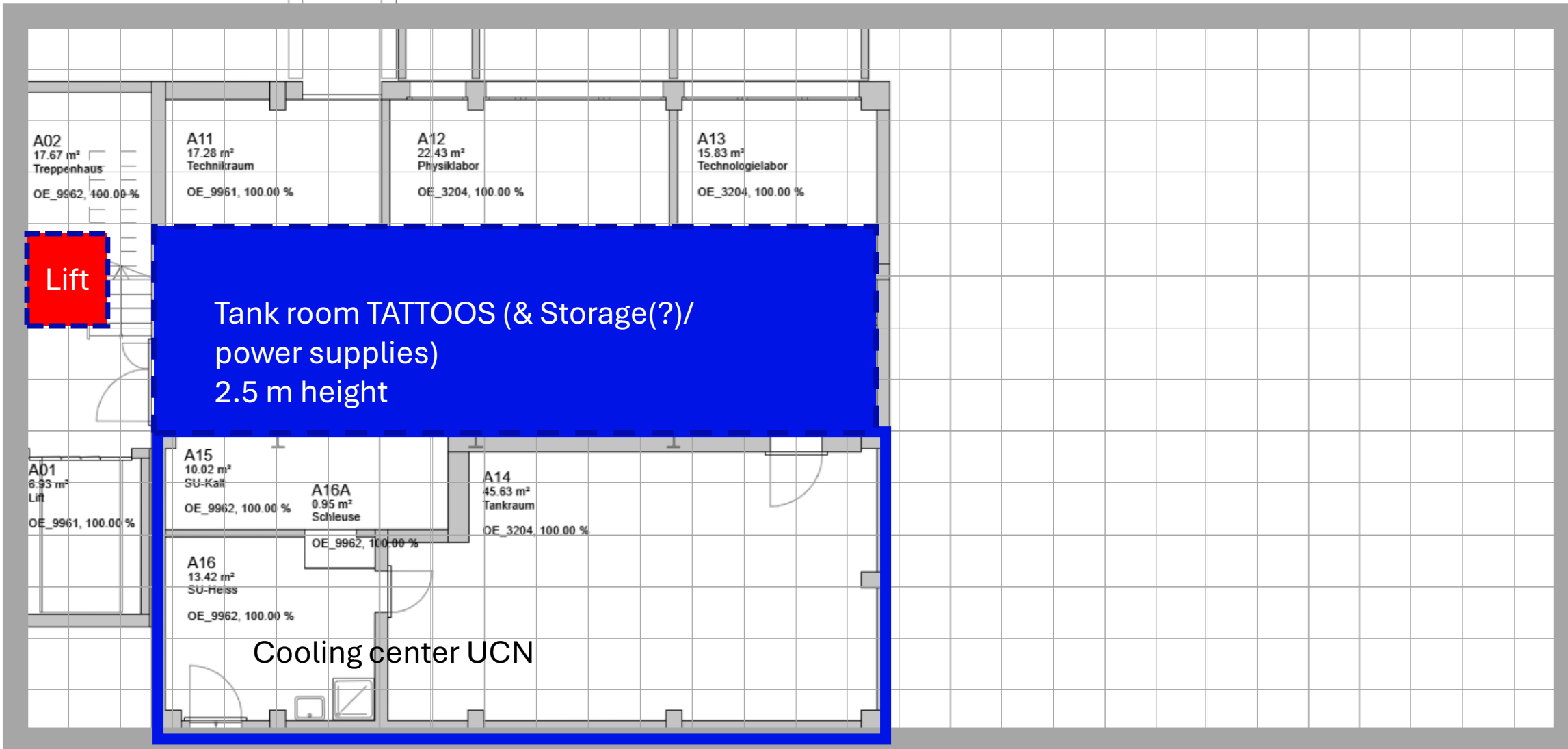
Ventilation & Filtering

Power supplies

Emergency door

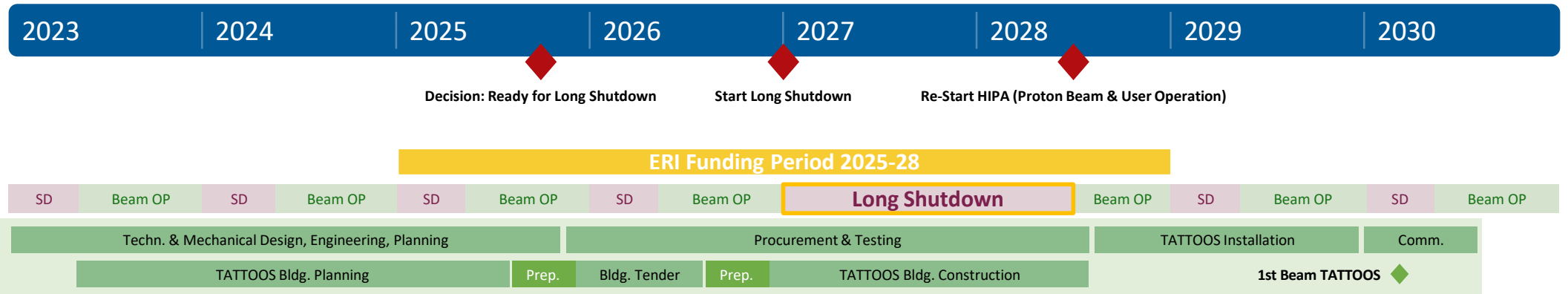
Stairs

Medienkanal



Cooling center UCN

Time-line TATTOOS



TATTOOS Collaboration

R. Eichler^{1,2*}, N. van der Meulen^{1,3}, D. Kiselev⁴, C. Baumgarten⁴, A. Ivanov⁴,
S. Jollet⁴, M. Hartmann⁴, S. Harzmann⁴, D. Laube⁴, M. Mostamand³, N. Preiss⁴,
T. Rauber⁴, D. Reggiani⁴, J. Snuverink⁴, S. Warren⁴, H. Zhang⁴

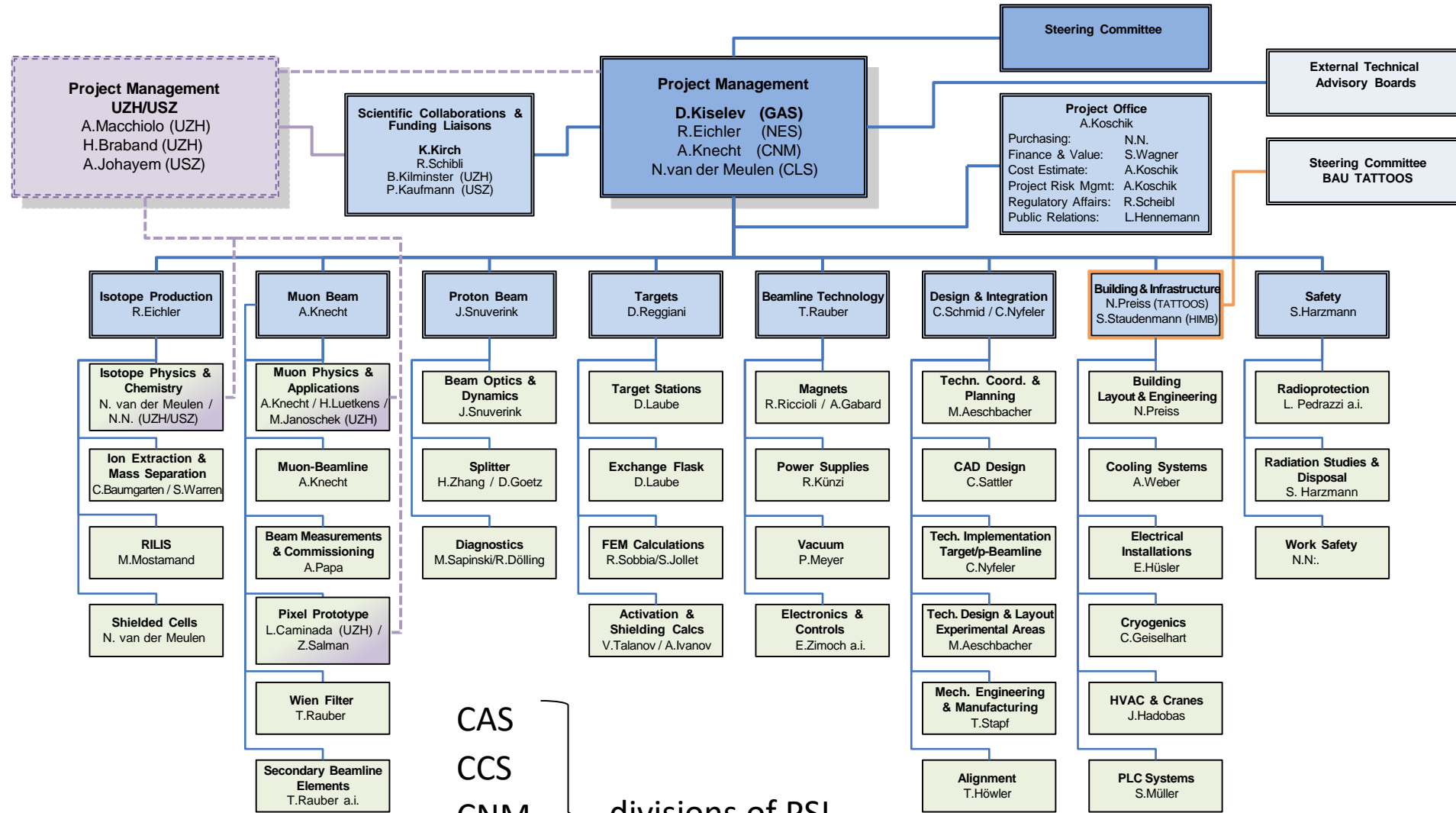
¹ Laboratory of Radiochemistry, Paul Scherrer Institute, Villigen PSI, Switzerland

² Department of Chemistry, Biochemistry and Pharmacy, University of Bern, Bern, Switzerland

³ Center for Radiopharmaceutical Sciences PSI-ETHZ, Paul Scherrer Institute, Villigen-PSI, Switzerland

⁴ Center for Accelerator Science and Engineering, Paul Scherrer Institute, Villigen PSI, Switzerland

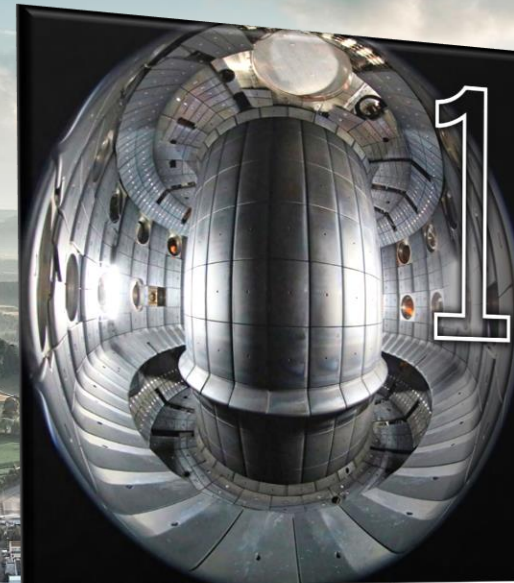
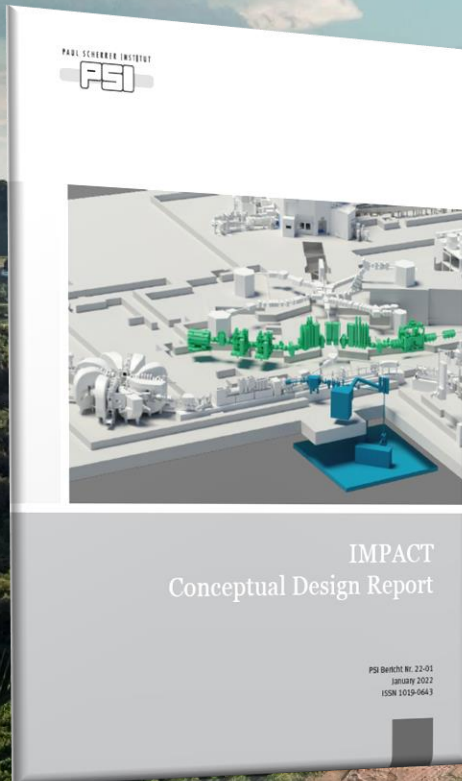
Organigram "People of IMPACT"



CAS
 CCS
 CNM
 CLS
 NES

divisions of PSI
 5 out of 8 divisions contribute to IMPACT!

TATTOOS as Part of the IMPACT Proposal is on the Roadmap of Swiss Research Infrastructures for 2025-28



Swiss Roadmap for Research Infrastructures in view of the
2025-2028 ERI Dispatch
(Roadmap for Research Infrastructures 2023)

Part I: National Research infrastructures

<https://www.psi.ch/en/impact>

https://www.sbfi.admin.ch/dam/sbfi/en/dokumente/2023/06/roadmap_forschungsinfrastrukturen_2023_teil_1.pdf.download.pdf/Roadmap_Forschungsinfrastrukturen_2023_Teil_1_EN.pdf

Calculated Production Rates at TATTOOS



Radionuclide Produced				CERN 2uA 600MeV	TATTOOS 100uA 590MeV*	Calculated patient doses
Element	Mass number	Target material	T _{1/2}	Irradiation time: 12h	Irradiation time: 6d	
					GBq**	
Tb	149	Ta	4.118 h	0.375	21	No reference
Tb	152	Ta	17.5h	0.038	5	No reference
Tb	155	Ta	5.32d	0.006	3	No reference
<i>Tm</i>	165	<i>Ta</i>	30.06 h	0.532	106	No reference
<i>Er</i>	165	<i>Ta</i>	10.36 h	0.662	60	No reference
Ac	225	UCx	9.92d	0.021	10	Ca. 1'300
Ra	225	UCx	14.9d	0.014	7	Ca. 700
Ac	225	ThCx	9.92d	0.024	12	Ca. 1'500
Ra	225	ThCx	14.9d	0.012	6	Ca. 800
<i>Rn</i>	211	<i>ThO2</i>	14.6h	0.139	12	Ca. 50
Ra	223	ThCx	11.43d	0.036	19	Ca. 4'500

* theoretical calculations; ** assuming ISOLDE yields